#### FIG. 1A

| cagg             | gato | ag ç  | ggtto | ccag | ga ac | ctcaç | ggato | tg0   | cagto | gagg | acca  | agaca | acc . | actga      | attgca | 60  |
|------------------|------|-------|-------|------|-------|-------|-------|-------|-------|------|-------|-------|-------|------------|--------|-----|
| gga              |      |       |       |      |       |       |       |       |       |      |       |       |       | tat<br>Tyr |        | 108 |
| gac<br>Asp       |      |       |       |      |       |       |       |       |       |      |       |       |       |            |        | 156 |
| cct Pro          |      |       |       |      |       |       |       |       |       |      |       |       |       |            |        | 204 |
| aga<br>Arg       |      |       |       |      |       |       |       |       |       |      |       |       |       |            |        | 252 |
| ggg<br>Gly       |      |       |       |      |       |       |       |       |       |      |       |       |       |            |        | 300 |
| cag<br>Gln<br>80 |      |       |       |      |       |       |       |       |       |      |       |       |       |            |        | 348 |
| gcc .<br>Ala     |      |       |       |      |       |       |       |       |       |      |       |       |       |            |        | 396 |
| ctt (<br>Leu     |      |       |       |      |       |       |       |       |       |      |       |       |       |            |        | 444 |
| Pro              |      |       |       |      |       |       |       |       |       |      |       |       |       | gcc<br>Ala |        | 492 |
| acc<br>Thr       |      |       |       |      |       |       |       |       | tag   | ggag | gacas | gga a | aact  | gcgti      | :t     | 542 |
| tago             | ctt  | gtg ( | cccc  | caaa | cc aa | agcto | catco | tg(   | ctca  | gggt | ctat  | ggta  | agg   | caga       | ataatg | 602 |
| tccc             | ccga | aa 1  | tatgt | cca  | ca to | cctaa | atcco | e aag | gatet | gtg  | cata  | atgti | ac    | cata       | catgtc | 662 |
| caaa             | gagg | gtt 1 | tgca  | aaat | gt ga | attat | tgtta | a agg | gatct | tga  | aat   | gagga | aga   | caat       | ctggg  | 722 |
| ttat             | cctt | gt g  | gggct | cagi | t ta  | aatca | acaaç | gaag  | ggagg | gcag | gaag  | gggag | gag   | tcaga      | agagag | 782 |
| aatg             | gaag | gat a | accat | gcti | c ta  | atti  | ttgaa | a gat | ggag  | gtga | ggg   | gcctt | ga    | gccaa      | acaaat | 842 |
| gcag             | gtgt | tt 1  | ttaga | aggt | g ga  | aaaa  | gccaa | a ggg | gaac  | ggat | tct   | cctc  | ag .  | agtc       | ccgga  | 902 |

#### FIG. 1B

aggaacacag ctcttgacac atggatttca gctcagtgac acccatttca gacttctgac 962 ctccacaact ataaaataat aaacttgtgt tattgtaaac ctctaaaaaa aaaaaaaa 1020

#### FIG. 2A

| cagg | gato  | ag g  | ggtto | ccag  | ga ac | ctcag | ggato      | tg:  | cagto | gagg | acca  | agaca | acc a | actga             | attgca | 60  |
|------|-------|-------|-------|-------|-------|-------|------------|------|-------|------|-------|-------|-------|-------------------|--------|-----|
|      |       |       |       |       |       |       |            |      |       |      |       |       |       | tat<br>Tyr        |        | 108 |
|      |       |       |       |       |       |       |            |      |       |      |       |       |       | gga<br>Gly<br>30  |        | 156 |
|      |       |       |       |       |       |       |            |      |       |      |       |       |       | cct<br>Pro        |        | 204 |
|      |       |       |       |       |       |       |            |      |       |      |       |       |       | cag<br>Gln        |        | 252 |
|      |       |       |       |       |       |       |            |      |       |      |       |       |       | tcc<br>Ser        |        | 300 |
|      |       |       |       |       |       |       |            |      |       |      |       |       |       | gaa<br>Glu        |        | 348 |
|      |       |       |       |       |       |       |            |      |       |      |       |       |       | ttc<br>Phe<br>110 |        | 396 |
|      |       |       |       |       |       |       |            |      |       |      |       |       |       | gca<br>Ala        |        | 444 |
|      |       |       |       |       |       |       |            |      |       |      |       |       |       | gcc<br>Ala        |        | 492 |
|      |       |       |       |       |       |       | agc<br>Ser |      | tag   | gga  | gacas | gga a | aact  | gegt              | it     | 542 |
| tago | ctt   | gtg ( | cccc  | caaa  | cc aa | agct  | catco      | tg:  | ctcag | gggt | ctai  | ggt   | agg ( | caga              | ataatg | 602 |
| tccc | ccga  | aa t  | tatgi | cca   | ca to | ccta  | atcc       | aa   | gatct | gtg  | cata  | atgt  | tac   | cata              | catgtc | 662 |
| caaa | ıgagg | gtt t | ttgc  | aaat  | gt ga | attai | tgtta      | a ag | gatct | tga  | aat   | gagg  | aga   | caat              | cctggg | 722 |
| ttat | cctt  | gt    | gggct | cag   | tt ta | aatca | acaa       | g aa | ggagg | gcag | gaag  | ggga  | gag   | tcag              | agagag | 782 |
| aatg | gaag  | gat a | accat | tgct  | tc ta | att   | ttgaa      | a ga | tggag | gtga | 9999  | geet  | tga 🤄 | gcca              | acaaat | 842 |
| gcag | gtgt  | tt 1  | ttaga | aaggi | tg ga | aaaa  | gccaa      | a gg | gaac  | ggat | tct   | cctc  | tag   | agtc              | ccgga  | 902 |

#### FIG. 2B

aggaacacag ctcttgacac atggatttca gctcagtgac acccatttca gacttctgac 962 ctccacaact ataaaataat aaacttgtgt tattgtaaac ctctaaaaaa aaaaaaaa 1020

#### FIG. 3

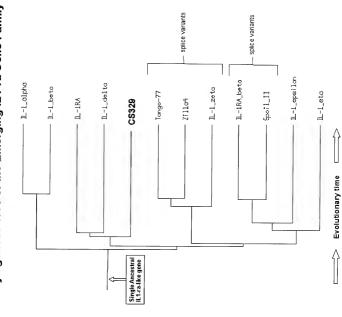
| gct              | cccg              | cca ç            | ggaga             | aaag             | ga ad            | catto             | tgag             | 999               | gagto            | ctac             | acc               | etgt             | gga g             | gctca            | aag              | 57  |
|------------------|-------------------|------------------|-------------------|------------------|------------------|-------------------|------------------|-------------------|------------------|------------------|-------------------|------------------|-------------------|------------------|------------------|-----|
| atg<br>Met<br>1  | gtc<br>Val        | ctg<br>Leu       | agt<br>Ser        | ggg<br>Gly<br>5  | gcg<br>Ala       | ctg<br>Leu        | tgc<br>Cys       | ttc<br>Phe        | cgt<br>Arg<br>10 | gag<br>Glu       | gac<br>Asp        | cag<br>Gln       | aca<br>Thr        | cca<br>Pro<br>15 | ctg<br>Leu       | 105 |
| att<br>Ile       | gca<br>Ala        | gga<br>Gly       | atg<br>Met<br>20  | tgt<br>Cys       | tcc<br>Ser       | ctc<br>Leu        | ccc<br>Pro       | atg<br>Met<br>25  | gca<br>Ala       | aga<br>Arg       | tac<br>Tyr        | tac<br>Tyr       | ata<br>Ile<br>30  | att<br>Ile       | aaa<br>Lys       | 153 |
| tat<br>Tyr       | gca<br>Ala        | gac<br>Asp<br>35 | cag<br>Gln        | aag<br>Lys       | gct<br>Ala       | cta<br>Leu        | tac<br>Tyr<br>40 | aca<br>Thr        | aga<br>Arg       | gat<br>Asp       | ggc<br>Gly        | cag<br>Gln<br>45 | ctg<br>Leu        | ctg<br>Leu       | gtg<br>Val       | 201 |
| gga<br>Gly       | gat<br>Asp<br>50  | cct<br>Pro       | gtt<br>Val        | gca<br>Ala       | gac<br>Asp       | aac<br>Asn<br>55  | tgc<br>Cys       | tgt<br>Cys        | gca<br>Ala       | gag<br>Glu       | aag<br>Lys<br>60  | atc<br>Ile       | tgc<br>Cys        | ata<br>Ile       | ctt<br>Leu       | 249 |
| cct<br>Pro<br>65 | aac<br>Asn        | aga<br>Arg       | ggc<br>Gly        | ttg<br>Leu       | gcc<br>Ala<br>70 | cgc<br>Arg        | acc<br>Thr       | aag<br>Lys        | gtc<br>Val       | ccc<br>Pro<br>75 | att<br>Ile        | ttc<br>Phe       | ctg<br>Leu        | ggg<br>Gly       | atc<br>Ile<br>80 | 297 |
| cag<br>Gln       | gga<br>Gly        | 999<br>Gly       | agc<br>Ser        | cgc<br>Arg<br>85 | tgc<br>Cys       | ctg<br>Leu        | gca<br>Ala       | tgt<br>Cys        | gtg<br>Val<br>90 | gag<br>Glu       | aca<br>Thr        | gaa<br>Glu       | gag<br>Glu        | ggg<br>Gly<br>95 | cct<br>Pro       | 345 |
| tcc<br>Ser       | cta<br>Leu        | cag<br>Gln       | ctg<br>Leu<br>100 | gag<br>Glu       | gat<br>Asp       | gtg<br>Val        | aac<br>Asn       | att<br>Ile<br>105 | gag<br>Glu       | gaa<br>Glu       | ctg<br>Leu        | tac<br>Tyr       | aaa<br>Lys<br>110 | ggt<br>Gly       | ggt<br>Gly       | 393 |
|                  | gag<br>Glu        |                  |                   |                  |                  |                   |                  |                   |                  |                  |                   |                  |                   |                  |                  | 441 |
| ttc<br>Phe       | agg<br>Arg<br>130 | ctt<br>Leu       | gag<br>Glu        | gct<br>Ala       | gct<br>Ala       | gcc<br>Ala<br>135 | tgg<br>Trp       | cct<br>Pro        | ggc<br>Gly       | tgg<br>Trp       | ttc<br>Phe<br>140 | ctg<br>Leu       | tgt<br>Cys        | ggc<br>Gly       | ccg<br>Pro       | 489 |
|                  | gag<br>Glu        |                  |                   |                  |                  |                   |                  |                   |                  |                  |                   |                  |                   |                  |                  | 537 |
|                  | cgt<br>Arg        |                  |                   |                  |                  |                   |                  |                   |                  |                  | tag               | gga              | gaca              | gga              |                  | 583 |
| aac              | tgcg              | ttt              | tagc              | cttg             | tg c             | cccc              | aaac             | c aa              | gctc             | atcc             | tgc               | tcag             | ggt               | ctat             | ggtagg           | 643 |
| cag              | aata              | atg              | tccc              | ccga             | aa t             | atgt              | ccac             | a tc              | ctaa             | tccc             | aag               | atct             | gtg               | cata             | tgttac           | 703 |
| cat              | acat              | gtc              | caaa              | gagg             | tt t             | tgca              | aatg             | t ga              | ttat             | gtta             | a                 |                  |                   |                  |                  | 744 |

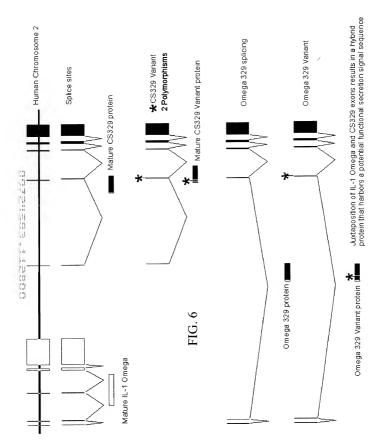
#### FIG. 4A

|                          | 1             |              |               |                    | 50             |
|--------------------------|---------------|--------------|---------------|--------------------|----------------|
| IL-1_alpha               | MAEVPKLASE    | MMAYYSGNED   | DLFFEADGPK    | QMKCSFQDLD         | LCPLDGGIQI     |
| IL-1_beta                | ~~~~~~~       | ~~~~~~~      | ~~~~~~        | ~~~~~~~            | ~~~~~~         |
| IL-1RA                   | ~~~~~~        | ~~~~~~~      | ~~~~~~        | ~~~~~~~            | ~~~~~~         |
| IL-1_delta               | ~~~~~~~       | ~~~~~~~      | ~~~~~~~       | ~~~~~~~            | ~~~~~~         |
| CS329                    | ~~~~~~~       | ~~~~~~~      | ~~~~~~~       | ~~~~~~~~           | ~~~~~~         |
| Tango-77                 | ~~~~~~~       | ~~~~~~~      | ~~~~~~~       | ~~~~~~~            | ~~~~~~         |
| Zilla4                   | ~~~~~~~~      | ~~~~~~~      | ~~~~~~~       |                    | ~~~~~~         |
| IL-1 zeta                | ~~~~~~~       | ~~~~~~~      | ~~~~~~~       | ~~~~~~~            | ~~~~~~         |
| IL-1RA beta              | ~~~~~~        | ~~~~~~~~     | ~~~~~~~       | ~~~~~~~            | ~~~~~~         |
| Spoil II                 | ~~~~~~~~      | ~~~~~~~      | ~~~~~~~       | ~~~~~~~            | ~~~~~~~        |
| IL-1 epsilon             | ~~~~~~~       | ~~~~~~~~~    | ~~~~~~~       | ~~~~~~~~           | ~~~~~~~        |
| IL-1 eta                 | ~~~~~~~       | ~~~~~~~      | ~~~~~~~       | ~~~~~~             | ~~~~~~         |
| _                        |               |              |               |                    |                |
|                          | 51            |              |               |                    | 100            |
| IL-1 alpha               | RISDHHYSKG    | FRQAASVVVA   | MDKLRKMLVP    | ${\tt CPQTFQENDL}$ | STFFPFIFE      |
| IL-1 beta                | ~~~~~~~       |              | ~~~~~~        | ~~~~~~~~~          | ~~~~~~~        |
| TT1PA                    | ~~~~~~~~      | ~~~~~~~      | ~~~~~~~~      | ~~~~MEIC           | RGLRSHLITI     |
| TI-1 delta               | ~~~~~~~       | ~~~~~~~      | ~~~~~~~       | ~~~~~~~            | ~~~~~~~        |
| CS329                    | ~~~~~~~       | ~~~~~~~      | ~~~~~~~       | ~~~~~~~            | ~~~~~~~        |
|                          |               |              |               | KDEPQCCLED         |                |
|                          |               |              |               | KDEPQCCLED         |                |
| TI1 zeta                 | ~~~~~~~~      | ~~~~~~~~     | ~~~~~~~       |                    | MSGCDRRETE     |
| IL-1RA beta              | ~~~~~~~       | MRGTPGDADG   | GGRAVYOS      |                    |                |
| Spoil II                 | ~~~~~~~~      | MRGTPGDADG   | GGRAVYOSSE    | SNAVGMGLWR         | LRPSALTLS      |
| IL-1 epsilon             | ~~~~~~~       | ~~~~~~~      | ~~~~~~~       | ~~~~~~~            | ~~~~~~~        |
| TL-1 eta                 | ~~~~~~~~      | ~~~~~~~      | ~~~~~~~~      | ~~~~~~~            | ~~~~~~         |
| 11 1_000                 |               |              |               |                    |                |
|                          | 101           |              |               |                    | 150            |
| II1 alpha                |               | FAYVHDAPVR   | SLNCTLRDSO    | QKSLVMSGPY         |                |
| II-1 beta                | 22222222      | ~~~~~APVR    | SLNCTLRDSO    | QKSLVMSGPY         | ELKALHLOGO     |
| TI.=1PA                  | LIFLEHSETT    | CRESCRESSE   | TOAFRIWDVN    | QKTFYLRNN.         | OLVAGYLOGE     |
|                          |               |              |               | LKVLYLHNN.         |                |
|                          |               |              |               | QKALYTRDG.         |                |
|                          |               |              |               |                    |                |
|                          |               |              |               | HKVLVLDSG.         |                |
|                          |               |              |               | HKVLVLDSG.         |                |
| IL-1RA_beta              | THOIGHOT HIGH | MCK          | DITCTINDIN    | OOVWILOGO          | NI.VA VPRS     |
|                          |               |              |               | QQVWTLQGQ.         |                |
| IL-1 epsilon             |               |              |               |                    |                |
| TI1 eta                  |               | MNDORFAA     | PKSYATRDSR    | QMVWVLSGN.         | SLIA APLS      |
| 111-1_000                |               | THE QUELTE   | I NO IIII DON | QI.T.T. DOGIT.     | 022111 1112 20 |
|                          | 151           |              |               |                    | 200            |
| TL-1 alpha               |               | SEVOCEE      | SMUKTOVALG    | LKEKNLYLSC         |                |
| II-1_alpha               | DMEQQVVFSM    | SEVOGEE      | SNDKIPVALG    | LKEKNLYLSC         | VI.KDDK PI     |
|                          |               |              |               | IHGGKMCLSC         |                |
| TI1 delta                | KVIKGERISV    | VDNDWI.DASI. | SPVILG        | VQGGSQCLSC         | GVGOE PI       |
|                          |               |              |               | IQGGSRCLAC         |                |
|                          |               |              |               | VSKGEFCLYC         |                |
|                          |               |              |               | VSKGEFCLYC         |                |
|                          |               |              |               | VSKGEFCLYC         |                |
| IL-1_zeta<br>IL-1RA beta |               |              |               |                    |                |
|                          |               |              |               | IQNPEMCLYC         |                |
| IL-1 epsilon             |               |              |               |                    |                |
|                          |               |              |               | TRGEDICTEC         |                |
|                          |               |              |               |                    |                |

#### FIG. 4B

|              | 201         |            |              |            | 250        |
|--------------|-------------|------------|--------------|------------|------------|
| IL-1 alpha   | LOLESVDPKN  | YPKKKMEK   | RFVFNKIEIN   | NKLEFESAQF | PNWYISTSQA |
| IL-1 beta    | LOLESVDPKN  | YPKKKMEK   | RFVFNKIEIN   | NKLEFESAQF | PNWYISTSQA |
| TL-1RA       | LOLEAVNITD  | LSENRKQDKR | .FAFIRSDSG   | PTTSFESAAC | PGWFLCTAME |
| TL-1 delta   | LTLEPVNIME  | LYLGAKESKS | .FTFYRRDMG   | LTSSFESAAY | PGWFLCTVPE |
|              | LOLEDVNIEE  | LYKGGEEATR | .FTFFQSSSG   | SAFRLEAAAW | PGWFLCGPAE |
| Tango-77     | LOLKKEKLMK  | LAAQKESARR | PFIFYRAQVG   | SWNMLESAAH | PGWFICTSCN |
| Zilla4       | LOLKKEKLMK  | LAAQKESARR | PFIFYRAQVG   | SWNMLESAAH | PGWFICTSCN |
| IL-1 zeta    | LQLKKEKLMK  | LAAQKESARR | PFIFYRAQVG   | SWNMLESAAH | PGWFICTSCN |
| IL-1RA beta  | LOLKEQKIMD  | LYGQPEPV.K | PFLFYRAKTG   | RTSTLESVAF | PDWFIA.SSK |
| Spoil II     | LOLKEOKIMD  | LYGOPEPV.K | PFLFYRAKTG   | RTSTLESVAF | PDWFIA.SSK |
| IL-1 ensilon | LOLKEKDIMD  | LYNOPEPV.K | SFLFYHSQSG   | RNSTFESVAF | PGWFIAVSSE |
| IL-1 eta     | LOLKEKNIMD  | LYVEKKAQ.K | PFLFFHNKEG   | STSVFQSVSY | PGWFIATSTT |
| _            |             |            |              |            |            |
|              | 251         |            |              | 290        | )          |
| IL-1 alpha   | ENMPVFL     | .GGTKGGQDI | TDFTMQFVSS   | ~~~~~~~    |            |
| IL-1 beta    | ENMPVFL     | .GGTKGGQDI | TDFTMQFVSS   | ~~~~~~~    |            |
| IL-1RA       | ADQPVSLTNM  | PDEGVMV    | TKFYFQEDE~   | ~~~~~~~    |            |
| IL-1 delta   | ADQPVRLTQL  | PENGGWNAPI | TDFYFQQCD~   | ~~~~~~~    |            |
| _CS329       | PQQPVQLTKE  | SEPSAR     | TKFYFEQSW~   | ~~~~~~~    |            |
| Tango-77     | CNEPVGVTDK  | FENRKH     |              | AEMSPSEVSD |            |
|              |             | FENRKH     |              |            |            |
| IL-1 zeta    | CNEPVGVTDK  | FENRKH     | IEFSFQPVCK   | AEMSPSEVSD |            |
| IL-1RA beta  | RDOPIILTSE  | LGKSYN     | TAFELNIND~   | ~~~~~~     |            |
| Spoil II     | RDOPIILTSE  | LGKSYN     | TAFELNIND~   | ~~~~~~~    |            |
| IL-1 epsilon | GGCPLILTQE  | LGKANT     | TDFGLTMLF~   | ~~~~~~~~   |            |
| TT 1 0#0     | SGQPIFLTKE  | DOTTININ   | TNEVLDSVE-   |            |            |
| IL-I CLA     | POOLITITION | TOT IIII   | INI I LLDOVE |            |            |





#### FIG. 7

|  | tgc<br>Cys        |  |  |  |     |  |  |  | 48  |
|--|-------------------|--|--|--|-----|--|--|--|-----|
|  | aag<br>Lys        |  |  |  |     |  |  |  | 96  |
|  | tca<br>Ser        |  |  |  |     |  |  |  | 144 |
|  | cta<br>Leu<br>50  |  |  |  |     |  |  |  | 192 |
|  | tgc<br>Cys        |  |  |  |     |  |  |  | 240 |
|  | gag<br>Glu        |  |  |  |     |  |  |  | 288 |
|  | cgt<br>Arg        |  |  |  |     |  |  |  | 336 |
|  | gct<br>Ala        |  |  |  |     |  |  |  | 384 |
|  | cag<br>Gln<br>130 |  |  |  |     |  |  |  | 432 |
|  | ttc<br>Phe        |  |  |  | taa |  |  |  | 459 |

#### FIG. 8

|     | MCSLPMARYYIIKYADQKALYTRDGQLLVGDPVADNCCABKICTLPNRGL |     |
|-----|--|-----|
| 1   | MCSLPMARYYIIKDAHQKALYTRNGQLLLGDPDSDNYSPEKVCILPNRGL | 50  |
| 51  | DRTKVPIFLGIQGGSRCLACVETEEGPSLQLEDVNIEELYKGGEEATRFT | 100 |
| 51  | DRSKVPIFLGMQGGSCCLACVKTREGPLLQLEDVNIEDLYKGGEQTTRFT | 100 |
|     |  |     |
| 101 | FFQSSSGSAFRLEAAAWPGWFLCGPAEPQQPVQLTKESEPSARTKFYFEQ | 150 |
|     | - 11 î.  |     |
| 101 | FFORSLGSAFRLEAAACPGWFLCGPAEPQQPVQLTKESEPSTHTEFYFEM | 150 |
|     |  |     |
| 151 | SW 152   |     |
| 131 |  |     |
| 151 | SR 152   |     |

#### FIG. 9A

actagtetee catagacaac agetgaatgt acgaggteag aageaaggee tgeeccagaa 60

ccattgcaag ccaggtgctg tcttgattgt agcctcataa aaaactgatg cagaattgcc 120 ccaccaacat gctccagatt cctgctccac agaaaccctg tgaactaacc atgttgcttt 180 tagattetge agtaagttga taatetgeag taaataacat tegatgaaag agaaacatgt 240 qtaqttactt tattatqatc aaaactttat ttctccactc tttccatttt ccttctcaga 300 attgacacca gcctttcact aacccaaata gcctatttaa atgctgatca tacttctctt 360 gttaactgtt acctgttccc aaaaggtaca attccctttc gaccatagct gcatctccca 420 cctqcacacc aggatqtttc tcatatttct acctaaaaca ttqqqqacta caaqtqaaaq 480 caaaagaggg ggtccatatc agaaccccag gtatttagct gtaaaactca cttgtcaggc 540 caqcttqaca qqtttacaqt ttqtaqaaqq accagaaaqa agqtaqccaa gacagaagag 600 qcaacctctq cttqtcctaq aaccttcaqt ccatatacat ctaaqctccc caqcaccatt 660 totaccacag acctotoaga gttcctgagg atgcagacco caggacactg acctoagttt 720 ccaqqcaqqq tttctqcaca ccccttcac actgcctgac tgggagttag tctcatggtg 780 caacactact ttgggacact gtacccatcc cctcgaccta cagaaaccat tcacttttca 840 aggtcacctc ctataggaag tatttgaaaa gatgagagtc atggtcattt gctatgataa 900 tattctqtgc ttatctccct qtaaaaagtt ggcttggggt ctctggcatg catctgacct 960 taaggttgga gctgcaccaa tatgttttta agcacccggc ataatgcttc gcaaaatttc 1020 agaacatggt ttgtacagaa tgtactttcc tccactcata caaacccttg taaaagagta 1080 qtttqaatcc caactcattc ttgaaggcca ccttttgtag ggtgacagaa tttaaaaata 1140 caqaatttaa aaatacttta tcccaqqqaa qctcacactt ctaaatccaq aatqaaaqaa 1200 gaaatagaaa cacacttgtg gtggcggtgg tggtggtgat ggtggtcgtg gtggtggtgg 1260 tqqtqqtqqt qqtqatqqtq gtqgtgqtgq tqqtqqtqgt ggtcqtgqtg gtqtaatgat 1320 cacagtaaag tgaggcatca tggcctgaga gagtcaggca tcacagctat tcaagtgaaa 1380 actacctact actgatttta gagttctata attttagtag cagccacagg cctggggcct 1440 qqqcctatat tttcagaqaq gaaatgttca caqcaqgtca actqcaqaca qtqaaqatca 1500 qaaatqtttc ataatcaqqt catcaqaqaa aaqqcaaaqq aqctqatqqa ctttatcctq 1560 aaaaagcaaa atccaaccca cctcatgctt aatgcattca aaggtctgcg ggcagaagaa 1620

#### FIG. 9B

tacattttgc tttttattat tataaattac ctggagaata tttttgtctg aattatctcc 1680

caaatattaa ccataaaaat aaaaaattcc atqtqtqctt ctcccaqqqq ctataaagcc 1740 cctggtctta gagttgttgg ggcaaaacct gacctttgaa gtagttactt ttgaagatgc 1800 cataccatac atttggccac ttggagagag tctaatgtca catctaaagg gttactctga 1860 tgctctgttt tctcatatgc ccttggctta cagctaacta tggctccagc taaactataa 1920 agtteettag caacagagat gatacqetat qtqtetttga cacagcagaa taaatgetta 1980 gtgaacatta ctgattgcct gacaggacac ctcacacttt ggtactttca acagagggat 2040 gtaaacttat gaagaacaat gaagaatgaa tattggcaat aaaagcaaaa attggttaac 2100 ccaattctag ctctgaaatc atttttaggt agtgggaagt ctttttgttt tgtttattca 2160 ctttacatcc caattgctgt cctccctcca agttccccac caccaccaca gtcctttttc 2220 cctccccttc tcctctgaga gaatggagaa ccctcctgga tattccccca tcatgaaaca 2280 trangictet graggetag acaetterer ragtgagger agtragggra gerragetag 2340 aaaaaqcata toocacaqac aqacaacaqc ttttqqqata qooccgttoc agttgtttag 2400 gatccacatg aaggctgagc tgcacatctg ctacatatga atgaggaggc ctaggtccag 2460 cctqtqtatq ttctttgqtt ggtggttcag actctgagag ccccaagggt ccaggtcagt 2520 tgactctqtt qqtcttcctq tqqacaccct qtccccttcc agcccacaat Ccttccccta 2580 atcettetee tteteaette cataaqaqtq tqaqqaqtet ttaaaaaacat gaagcatttt 2640 atctccccag ggcaacacat ggaaatgaaa gattgtgaaa agtaatttaa agaaaaagaa 2700 aaaaaaattt aacaaqqaat aaqaatcttg tttctctgaa aatgcttaag agtgtggaaa 2760 acataaactg gattctaata gaatgcaatt ggattgtaat gaaaacctat caaagttatg 2820 aaataqcttt cactaccttg cacaaaatct cttggcatgt gtgttgttgg caaattttct 2880 tqttaqttta aaaccacaac aataacaaca aaataqcaaa aattggqtct cagcctcatt 2940 cattttttct catttcttqc tctqtqatcq tctqqqtctt aaqctqacac ctcaccaatt 3000 cctcatcaaq acctttqtqq aaatttqcaa atgtcccaaa aaggagaatt acaataagtc 3060 agagaacqtt ctqtccaatt ctttatccct aqtqatqqat qaqtaaaqqa tqtataaqag 3120 atggataaat ggactgatgt acagataaat gaaggaatat gtacatggtt aggtggatag 3180 atqacttact caacaqatqa qtaqaaqqat qagaaataqa tgqacagctg gactgaggca 3240

#### FIG. 9C

tqcaaaqtca actqqaqaac tqaqtctctt gaccatgcac tqtccagggt ctcatattcc 3300 ctagagteca gggeceatgg etectgtgce atceccatge aaatetaagg ttaatacgtt 3360 ctacaqctqa qtttccttac atatgtgtct cagtaagttt gtatcaacta attaaatctg 3420 aaaggagtto ottotgatot toocaaacag agccacacto gtgatgaagt cagccotgot 3480 tcattqtgqt tctctggatg catctggctt ccatcagcat aatctttcta ttcttgatcc 3540 ttccaacctc ttcaggtctc agacagaacc ccatggagca tcaaagaggt ttgaccccag 3600 cattetttat etagetecaa aaccactaat aacacaetca ateacaetae ctacaeaac 3660 ageaggteag tgtctggcct ctgtcaaggc tttatgagtg actetctccc cttcccgcaa 3720 atactcatta atctccccac ctccttatta tttqqactqt qttqaaqata ttatqaaatc 3780 totgggotot tottocogga totagagoca attacagatt otgtaggttt gacccaccot 3840 qaccaqacat tataaacaca qtqctqqtqc cctqaaqaaa acaqttqqaq actccaqqca 3900 ttagaatcca ggcaccagga actacaggtc agtggtgaca gtcggtctct ctgtgtatct 3960 cttacacaca cacacataca cacacacaac acaacataca cacacataca acacacaaca 4020 catacacata caacacatac acacacaca cacttttctg taatgtctcc aaaattctca 4080 gqctctaggg aagaagaaat gtcttttaga gaatgcggtg tgatgttcta taagtctagg 4140 aatacttqat aqaatttaat qaqaaqtata gattaggtca aagcaagggt actacatatt 4200 tggaaccaca gagttttgaa agtcatctca aaagaaatta tttaggccag agatgttcaa 4260 aaaatgtttt gtttgtgaca tatggaagct cccatggaga cattctgtga ttctcatcaa 4320 tagacagtag ggatgccacc aaggtgctaa cgtcttcatc accccatcat ctatcataca 4380 tccaaatqqt ttctttqaaa acaatctcct tqtqaaactt aaaqtaqcct tqaaaatata 4440 ataatcttgt ccagcctctc atttcaatgg gaatagattg aaggcctaag gaccaaaaca 4500 taagttattt ttagaatcca gcctttcagt caaagcttga ttcatgcata tctgtgttct 4620 gatcttaagg tgctgtgtct gtcagttgta tagttggata gaggtacaga tgagctatat 4680 acatcatqct tcaaqatttc aqqatcttat aacttttata aaqcaaataa tttgtcttaa 4740 tgcacactaa taaacaatat agcaaagttt gacaggagtt cagagtactg ttagagaagt 4800 gaagggaaga attttgttat gatagtaaag gggaaaatca aattttgagt catggaatca 4860

#### FIG. 9D

tacatagttt gacatagaaa gaaccttggc aaccacataa totaatgcat gagcccaaga 4920 actggcctgt gtttttaaga tctcattctc agctgttatg taactgaaca gacaagatac 4980 taagcccaag tatagtgaag ccatgtccag tgatcttaat aggagtgaca ggaatggttg 5040 qtqatgaaga ggggtggatt ttgagcagga ataccaaaag caatgctgac tgtgcccttg 5100 gagagaatta gcatgagtcc ttgagagaaa aatgagatgc tattgcacaa gcaacctagg 5160 qccaqatggt gtcaagatag gtggccatcg tggactttag aaccaggcag gaatgtgatc 5220 agagatgtac tttatgtagg ttaggtttga ttcagaaacc aggagggtta gcatgtttac 5280 aatggtgact aaaaacaagc acaaggttat actttaaaga aataatctct gaaaagaagg 5340 gaggtatatt ttcagtgccg gaaagaggaa tattacaaaa gtgagaggag tagatttgag 5400 aaagagaagt ggattgtgga ggagcagatg ctcaccacgc ccttacactc acttgaactg 5460 acacccaaag atgaaggtgt gctgtggact gctgaagctc agcctgtggc tgggaagcag 5520 taaacaaaat tgctcatcac agctgtacaa gatattccat agcatataaa aataaaagtg 5580 cttaggctat tetettacaa eteteageet tatgaatgae eeggaaggaa aagaacteta 5640 caatgtgcct gtgtctgttc ttacttcctc tgccacaagc aaaagagcct tgggaattgg 5700 ctcagaggga acgtcatcaa acaggctggc cttgaggctg ggctgttatt cgtctacctg 5760 ggatagagga attogotatt ottttataat ocaagtgtgg ootggggaco agcagcatta 5820 ttaagacctg gttgcatgtt tgaaatgcag tctcagattt catcccagac ctaaagagta 5880 acactgtttt catgaggata caagattaag aaatatgcat tagagagtaa ttggctaaat 5940 gggtaaatgt catgcaagca ggaggatctg attgactccc caggacccac acagttccca 6000 tgccgtagag cacatctgta atcacagtag gcgtatgatg aaatgggagg tgaatcaaga 6060 gaatototag cagotacggg otggocagoo toccatgoac agcactaaat aaggoaagga 6120 ccaatacctg aagttgtccc attaccttca catatacacc acggcatgtg tgtacttgta 6180 ctcacacata caaacaaata cacacgtgca cacatacaaa actcagagat taaggacaat 6240 tggcctgaca tatcagttcc taagcctggc tcattgcttg taacactaca agcagtatta 6300 aataaggata ggcgagagaa cagttaccga atggttcaga agtggggcca tgcctgtgac 6360 tttaaacaaa tgtttcatat ttttaaataa taacacttag attacaaaat aaatttacta 6420 caggaaaatg ttaagaacta tcaacaacca ttgactatcc tgtcggccac aaatgagtgt 6480

#### FIG. 9E

tataacaage accageegte ettgtecaca tgtgtgtgtg tetacacage tatgaattta 6540 attgggataa taatgtgcac attctttacq gcctgcagtt tttacttcat qtatttgaaa 6600 tgtttgtgcc acaaatgtca tctttaagga gcatatcctt atttcctgga tttatcattc 6660 cctttcagcc gactggacat tgacagcatt tccaactttt caaccttgta aaaataacta 6720 attgaactat tttataacta agcatttggg caatcaatta cctctgcctg gaatgggggc 6780 aacaacacat gcaatcatgg gaaagccagg atgctgctgt ctgatcccta gccctqgcat 6840 togtgoagaa cotcactoto atotgtgooo tgatatoott cactotcaag tottttooca 6900 gtgactttta aaggcaacag aatcatatag ccaataatga aagctacttg gtctacagtt 6960 gtgtggcgtt ttttatagat attttcttca tttacatttc aaatgctatc ccaaaagtcc 7020 cetataccet eccecacet getecectae ecacteacte ceaettettg geeetggett 7080 tcccccttac tggggcatat aaagtttgct agaccaaggg gcctctcttc ccaatgatgg 7140 ccaactaggc cattttctgc tacatatgca gctagagaca ccagttctgg ggttactggt 7200 tagttcatat tgttgttcta cctatggggt tgcagacccc ttcagctctt gagtactttc 7260 tctagctcct ccattgggag ccctgtgttc catcctatag atgactgtga gcatccactt 7320 ctgtatttgc caggcactgg catatgaaat agtatctgca tttggtggct gattatggga 7380 tggacccccg ggtggggcag tetetggatg gtccateett teatettage tecaaacttt 7440 gtototgcaa ottottocat ggatatttta gtocotaato tagggagaaa tgaagtatoo 7500 acaagttgat cttccttctt gattttctta tgttttagaa gttgtatctt ggatattcta 7560 ggtttctggg ctaatatcca cttatcagtg agtacatatc aagtgaattc ttttgtgatt 7620 aggttacctc actcaagatg atattctcca ctatgttcat agcagcccta tttatagtag 7680 ccagaagetg gaaagaacce agtccctcaa cagaggaatg gatacagaaa atgtggcaca 7740 tttatgcaat ggagtaccac tcagatatta aaaacaacga atttatgaaa ttctcgggca 7800 aaaccctatc taaagaccag gaataaggaa aagatggact gcctgcctgc agctgggaga 7860 gctggggaga cctttgtgga ttctgtaata cttaggggta cggaacagct tgtggctgga 7920 taattctgag ctccagcatg tctgcccccc aaaaaacatt ctgtttttct gaaagccttt 7980 ttettetttg ceteagtgaa gaccagacae teecaactge agga <u>atq tgc tee ett</u> 8036 ccc atq qca aqa tac tac at gtaagtaa tettaacgat egeteaatca 8084

## FIG. 9F aggggcctgg agatcacatg agaagggaaa aggctgagtc aaagggacaa agctccctct 8144

agccacagaa atctcaaaca ctgaataatt gatcttcatc tttgtcaatc acaacagccc 8204 tctttcctgg tgacagaatg gaacaactgt aagagtggta ttgcttagtc cattttacag 8264 acceggaaac tcaacctcca cgaggttata caattttcct catgtcatgc aattacccaa 8324 aagcagagag tgggatcgga ctctctgttc tctaaactga tgtagctagt tcttagaaag 8384 ctcaaacaat cttgagtccc aaggacagca cctttatggt cacctggatt gatacctata 8444 tcaaaaaaaa aaaaaggtot cactagatag cootggotac cotgaaacto tcactgtgta 8504 catttaggtg accacgaact cacagagatc tgccttccaa gtgctgggat taaagtatgt 8564 accaccacac etgeatettt gacaataact gagtggtate taaattette cagtggetaa 8624 acagttaagt cccagttccc aaagtctgag aaaaatgcca ggtggtgaaa tctgtacaga 8684 cctttgttct taatgtacaa gtgagcctgc tttaaaaaaca atacgcaagc tgtttttgct 8744 attgctaagt gttgcagaga cagaaaaggc tcccagaagt ggtaactttg gtccagaggt 8804 totgttotca aactcattgt gagototgaa agcaactgat gggcagotot gaaatcagot 8864 gggcaattag gctaataaca ggcataattt taatgtttca cacgcatgac agttcctccc 8924 cagetgeect agtacatact taccetecta ggeaegteat tagacecata ggtataacca 8984 gtgactaatc aggccctggt ctaattctaa gttggcctcc tatataagtg ccactcagag 9044 tgtacctcat catggctgta gtgggcccag agtctaggga catagacttt tctattgtcc 9104 atcaaqctca qaaacatcat qqataqqqtt cattqtatct ccagggtacc tgagcttcaa 9284 agcaactcct cagacagcca tgaaaacatc ctcaattacc tcatgagaag acactattgt 9344 catttctqqa qcctctqata atcctgagcc taggcagctt tgggatgaaa caatttctac 9404 ccttattgga acagtgtccc tctcctgtct ggaaacaatt caccaaaggc tccatgtggt 9464 tgtccagtaa ggtggtatgg ggacagaaat ggacaatgat ccctgagggc agtgatccat 9524 taacettgee etectattte ag a ate aag gat gea eat caa aag get ttg tae 9577 aca cgg aat ggc cag ctc ctg ctg gga gac cct gat tca gac aat tat 9625 agt cca g gtgatcttc cggtggtggg ggtgggggag tggaggggag ggtgtggggg 9681

#### FIG. 9G

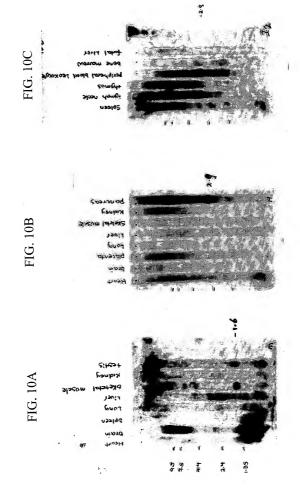
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|--|-------|
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| tggtcccctc actgtctgaa ttgcattctg tcttacctcc cag ag aag gtc tgt atc | 9978  |
| ctt cct aac cga ggc cta gac cgc tcc aag gtc ccc atc ttc ctg ggg    | 10026 |
| atg cag gga gga agt tgc tgc ctg gcg tgt gta aag aca aga gag gga    | 10074 |
| cct ctc ctg cag ctg gag gtgagacacc cctcctcatt gcagtcagta           | 10122 |
| ctgccactgg aacatagtga catctttgaa cccacatgtc ccctctcttg tttcccatct  | 10182 |
| atctctcttt gcctccagct gagggactct agcctttggg gatgtacaga aagaacatgg  | 10242 |
| cttcggaaaa ctcttcccta ttgagtcctt ctttggccaa gcctctgagg cactaagggc  | 10302 |
| tgacgtccca accaaacact catttcatct cacagctgtc tccctttccc cacag gat   | 10360 |
| gtg aac atc gag gac cta tac aag gga ggt gaa caa acc acc cgt ttc    | 10408 |
| ace ttt tte eag aga age ttg gga tet gee tte agg ett gag get get    | 10456 |
| gee tge eet gge tgg ttt ete tgt gge eea get gag eee eag eag eea    | 10504 |
| gtg cag etc ace aaa gag agt gaa eee tec ace cat act gaa tte tae    | 10552 |
| ttt gag atg agt cgg taa agagacataa ggctggggcc togtctagtg           | 10600 |
| ccccagtct gagatcttct tgctcagcat ctctggaaag cagaataagg aagataccaa   | 10660 |
| agatgtttgg gtcttaatcc ccagaatctg tgaccgtgtt acattaaatg gcaaagggat  | 10720 |
| ttttttttttc cttcatggtc catttgggcc cattggaatc atctgaggcc tcatgaggag | 10780 |
| aaggaagagg tcagagggag actggggcaa actttggtac taaaagtaac aatggagaca  | 10840 |
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| ttaagccact gagtttgaga tcattcaatg aagctgtcat aataaaacct gtgcttcaca  | 11020 |
| tacaattcaa tattggtagg caccccggtg atttcttgga aagacatcta gggattctcc  | 11080 |
| tggatgctga ttccagggtc cagtggagtc cctgggttga agagatttca caacccagaa  | 11140 |
| catcaggete gactetteta aaagteegte gttgcaccce ttgcetgaga gcattagcaa  | 11200 |

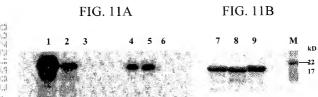
### FIG. 9H

ctgatttagt qaqqqtqaq ctgctqqcac ttttttqtqt caccagtqtc ttaagcagtg 11320 atggaggaca aaagatettt actgagaaga tggccatgaa getetggeta gacaccaaga 11380 atatqatata aqcaqaqcta caqcacaaqa tqaqccaatg aggaaagcca ttcagggagg 11440 ctaagcccag cttcccaaag ggacagctaa ccctggactc aaatgaatag gggttttcct 11500 qqcaqaqaac ataqqtcaaq cattctagqt agaatcagca attcagaaag gtgtgagaga 11560 qqcatqqaqa qctccaqqca tqtctqqqct atqqtqtqtc attcttqtqq caaqaatcca 11620 acqtctgtgq ttaaggagtt gctgaaaatt aaaataggaa aatgggtaga gtctaattgt 11680 qaatqacttq caaaqqaqtt taqcccataa qtqqqqaqct cagaqqaqtc atctaaqqat 11740 tgcaagcagg ggccctgtga tcattgctgg accagcctag gtgctacaga gcctaccttc 11800 aqctctqcat cctcactcac atccaggtac cttcagaggt caatttctgt gctctggttc 11860 tatqqqtaqc ctqaccctqt ttcatcttct tqtataactt aqqcacataa qcttaggqac 11920 tggtagagtt tacttgagtg attggtgaat caggcagcac caaactacaa gttgttcagg 11980 gctttaccaa gggggcactg attggagaat tggaatgagg gtggttagaa tgcattcaga 12040 aaacaagggg aagaaaaatt tgattgctta aagtggaaag tcccaactta aatgttagtc 12100 aqtaqtttct aattacttga gtctctaatt agaggttagt tggcagtttc tggttagtta 12160 atctaagttt cattttctta qqctatqacc attctctqaq tcqcatqtta qcaatqcaqt 12220 aagaactcaa gacccagaat agcctctgtt aattatttta gcaatgatca ctcatttctg 12280 ttqcctccta ttqaqatctq ttcccatqqa ccacccaqqc acatcaqqcc tcctagtacc 12340 aacataataa tgattgctgc acagacaaaa tatttttttt cagtatctgg tatttgctac 12400 atttccatta qtqctqqagq qaaqqctaca acqaccatga aggcatggcc cctgccttct 12460 aaqqacttac aatqtaataq qaqccctqac attataaaqt qqqtcacctt qtttcaaact 12520 gagccaaact gaggctgagg gcttagatta gtggtaggtc actttccaga catgttcagt 12580 gctaaqaaaa acacattctg gggttagtta gatgttttag ttcatttgat aagaagccca 12640 atgattggac tttcaacttc tggaacccat gtggtggaag agagaaccaa cttctgacca 12700 tttqqqtcat qqcacatccc ctaccatcac aaqaactcac caaaataaat taqaaaaatc 12760 aaqaaaaact catatcctat agacctctgg tagaattagc agaacgctgc tgtggcactt 12820

#### FIG. 9I

gggatttgaa actcaaaaat ggaagaagct acttgtgacc gttcaagact ccagggagc 12880
tcctctgaca catcccacga ctcaggctta aattccttct tctccctaga aggccacgcc 12940
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# 

## FIG. 12

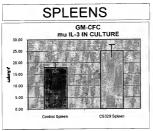
## Spleen

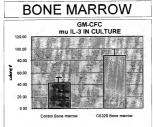
|              | CD4+  | CD8+ | CD4+ CD8+ | CD3+  | NK1.1+ | CD3+ NK1.1+ |
|--------------|-------|------|-----------|-------|--------|-------------|
| control mice | 20.18 | 3.72 | 1.67      | 24.07 | 3.06   | 1.4         |
| CS329 mice   | 15.89 | 3.99 | 0.37      | 22.9  | 2.08   | 1:1         |
| Difference:  | 4.29  | 0.27 | -1.3      | -1.17 | -0.98  | -0.3        |

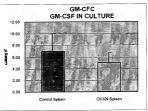
## **Bone Marrow**

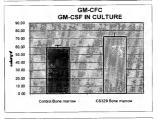
|              | CD4+ | CD8+ | CD4+ CD8+ | CD3+ | NK1.1+ | CD3+ / NK1.1+ |
|--------------|------|------|-----------|------|--------|---------------|
| control mice | 2.62 | 2.54 | 0.49      | 3.88 | 1.26   | 0.49          |
| CS329 mice   | 2.46 | 2.35 | 0.41      | 4.42 | 1.53   | 0.57          |
| Difference:  | 0.16 | 0.19 | 0.08      | 0.54 | 0.27   | 0.08          |

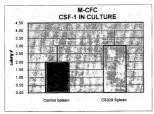
FIG. 13A











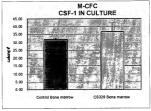
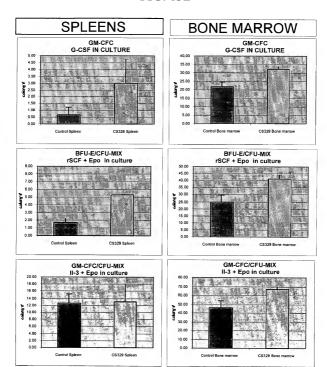


FIG. 13B



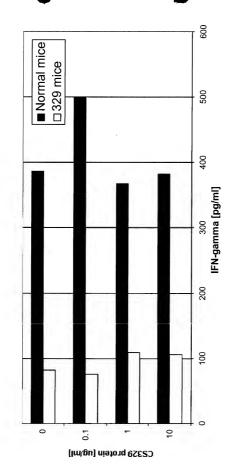


FIG. 15

